

TABLE 1
50% INHIBITORY CONCENTRATIONS IC₅₀
FOR FORMULA A

Example	R ¹	A	mICE IC ₅₀ (μM)	CPP32 IC ₅₀ (μM)
4	CH ₃	Ala	0.177	>10
7	CH ₃	Pro	11.7	>50
10	CH ₃	Val	0.531	2.48
13	CH ₃	Leu	5.52	5.62
16	CH ₃	Phe	3.34	49.8
21	CH ₃	Gly	34.7	>50
24	CH ₂ Ph	Ala	0.393	>50
27	(CH ₂) ₂ CH=CH ₂	Val	0.313	1.45
30	CH ₂ CO ₂ H	Ala	1.63	>50
33	(CH ₂) ₂ CO ₂ H	Ala	0.198	>50
reference	--	--	0.064	47.0

Fig. 1

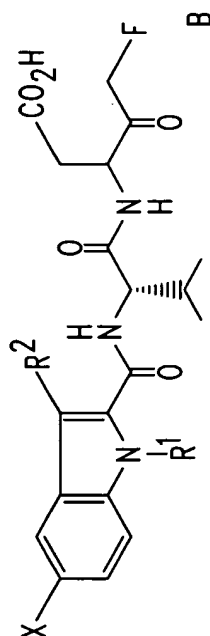


TABLE 2 DISSOCIATION CONSTANT K_i AND INACTIVATION RATE k_3/K_i FOR FORMULA B							
Example	R^1	R^2	X	mICE K_i (μM) k_3/K_i ($\text{M}^{-1}\text{s}^{-1}$)	CPP32 K_i (μM) k_3/K_i ($\text{M}^{-1}\text{s}^{-1}$)	Mch2 K_i (μM) k_3/K_i ($\text{M}^{-1}\text{s}^{-1}$)	Mch5 K_i (μM) k_3/K_i ($\text{M}^{-1}\text{s}^{-1}$)
43	CH ₃	CH ₃	H	1.40 2,860	0.960 13,400	0.017 58,800	0.062 21,500
46	CH ₃	Cl	H	1.68 6,150	0.830 25,900	ND	0.099 37,000
49	CH ₃	Cl	F	1.10 7,120	0.493 72,700	0.014 71,400	0.054 52,500
52	(CH ₂) ₃ Ph	H	H	0.133 45,100	0.742 33,700	0.024 41,700	0.077 32,500
55	Ph	H	H	0.843 8,900	0.110 74,200	0.036 55,600	0.043 35,300
58	CH ₂ CO ₂ H	H	H	0.327 16,800	0.125 58,700	0.051 19,600	0.038 127,000
61	CH ₃	H	H	0.240 41,700	0.520 21,200	0.033 30,300	0.026 38,500
62	CH ₃	CH ₃	F	0.397 7,560	0.113 44,200	0.040 25,000	0.102 29,400
63	(CH ₂) ₂ CH=CH ₂	H	H	0.327 18,300	0.125 56,000	0.104 19,200	0.038 131,600
64	CH ₃	H	F	0.234 21,400	0.180 38,900	0.052 38,500	0.063 47,600
65	CH ₃	CH ₂ CH(CH ₃) ₂	H	4.56 1,540	2.28 7,910	0.023 43,500	0.063 31,700
66	CH ₃	(CH ₂) ₂ Ph	H	0.632 14,200	0.505 21,800	0.038 26,300	0.051 39,200
67	CH ₃	H	OCH ₂ Ph	0.739 14,900	0.346 31,800	0.040 25,000	0.062 16,100
reference	--	--	--	0.015 278,000	0.820 14,600	0.594 3,370	0.018 83,300

Fig. 2

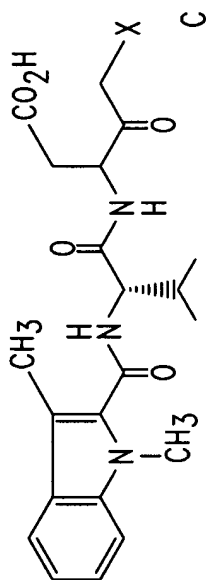


TABLE 3
 DISSOCIATION CONSTANT K_i AND INACTIVATION RATE
 k_3/K_i FOR FORMULA C

Example	X	mICE K_i (μM) k_3/K_i ($M^{-1}s^{-1}$)	CPP32 K_i (μM) k_3/K_i ($M^{-1}s^{-1}$)	Mch2 K_i (μM) k_3/K_i ($M^{-1}s^{-1}$)	Mch5 K_i (μM) k_3/K_i ($M^{-1}s^{-1}$)
43	F	1.40	0.960	0.017	0.062
70	OCO (2, 6-di-Cl-C ₆ H ₃)	1.16	0.052	0.030	0.364
71	OPO (C ₆ H ₅) ₂	0.124	0.046	0.060	0.022
72	O (1-Ph-3-CF ₃ -pyrazol-5-yl)	0.873	0.300	0.050	1.39
73	O 3-CONH ₂ -2-naphthyl)	8.00	1.58	0.632	0.213
74	O (2-CONH ₂ -1-phenyl)	0.297	0.419	0.340	0.547
75	OPO (CH ₃) ₂	4.33	1.05	ND	0.663
reference	--	0.015	0.820	0.594	0.018
					21,500
					2,750
					45,500
					720
					0
					0
					1,510
					83,300

Fig. 3

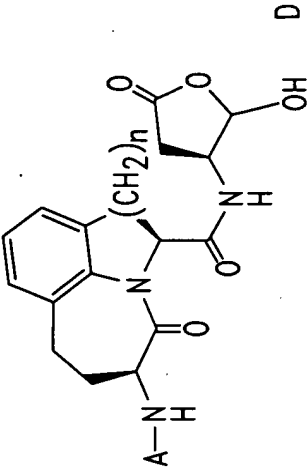
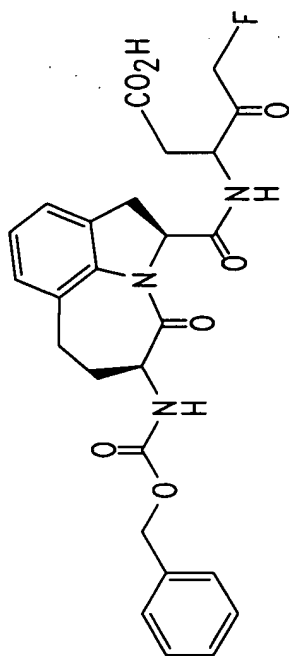


TABLE 4 50% INHIBITORY CONCENTRATIONS IC ₅₀ FOR FORMULA D						
Example No.	A	n	mICE IC ₅₀ (μM)	CPP32 IC ₅₀ (μM)	MCH-2 IC ₅₀ (μM)	MCH-3 IC ₅₀ (μM)
78	Cbz	1	0.019	1.03	40.1	6.96
82	Ac-Asp	1	0.694	0.0014	6.47	0.145
85	succinyl	1	0.571	0.245	1.81	2.83
88	Cbz-Asp	1	0.096	0.00052	ND	0.084
91	dihydrocinnamoyl	1	0.045	0.780	>10	32.6
94	Ac	1	3.07	3.87	>10	>50
100	Benzoyl	1	0.159	8.77	>50	>50
97	1-Naphthoyl	1	0.010	2.91	>50	12.3
103	Cbz	2	0.026	0.437	32.0	1.11
reference	-	-	0.064	47.0	>10	>10

Fig. 4



Example 106

TABLE 5 DISSOCIATION CONSTANT K_i AND INACTIVATION RATE k_3/K_i FOR EXAMPLE 106			
Enzyme	Example 106		Reference
	K_i (μM)	k_3/K_i ($M^{-1}s^{-1}$)	K_i (μM) k_3/K_i ($M^{-1}s^{-1}$)
mICE	0.0005	12,000,000	0.015 214,000
CPP32	0.012	960,000	0.820 12,200
MCH-2	0.033	25,000	0.594 2,950
MCH-5	0.022	98,000	0.018 83,300

Fig. 5

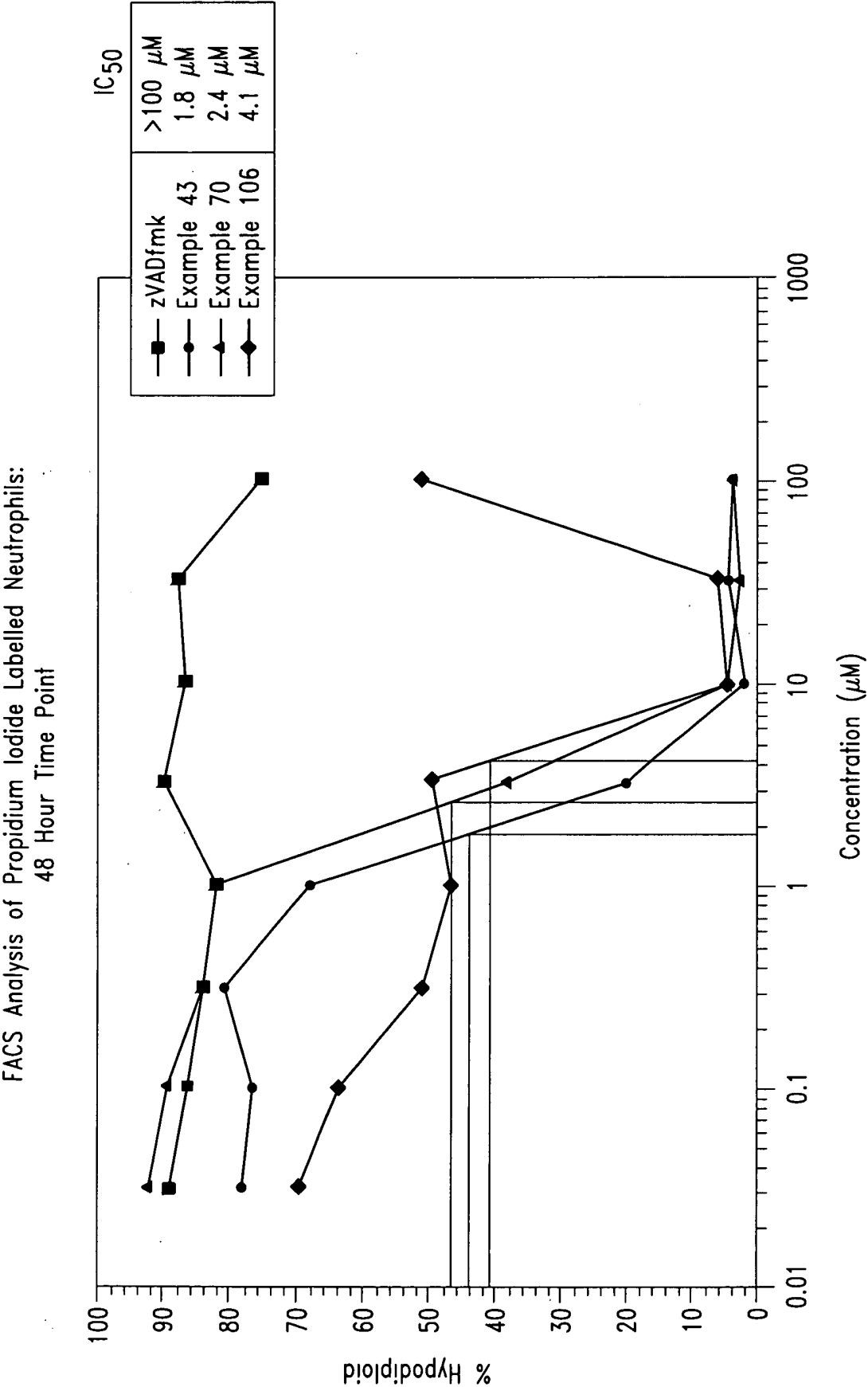


Fig. 6

Neutrophil Survival and Burst Assay

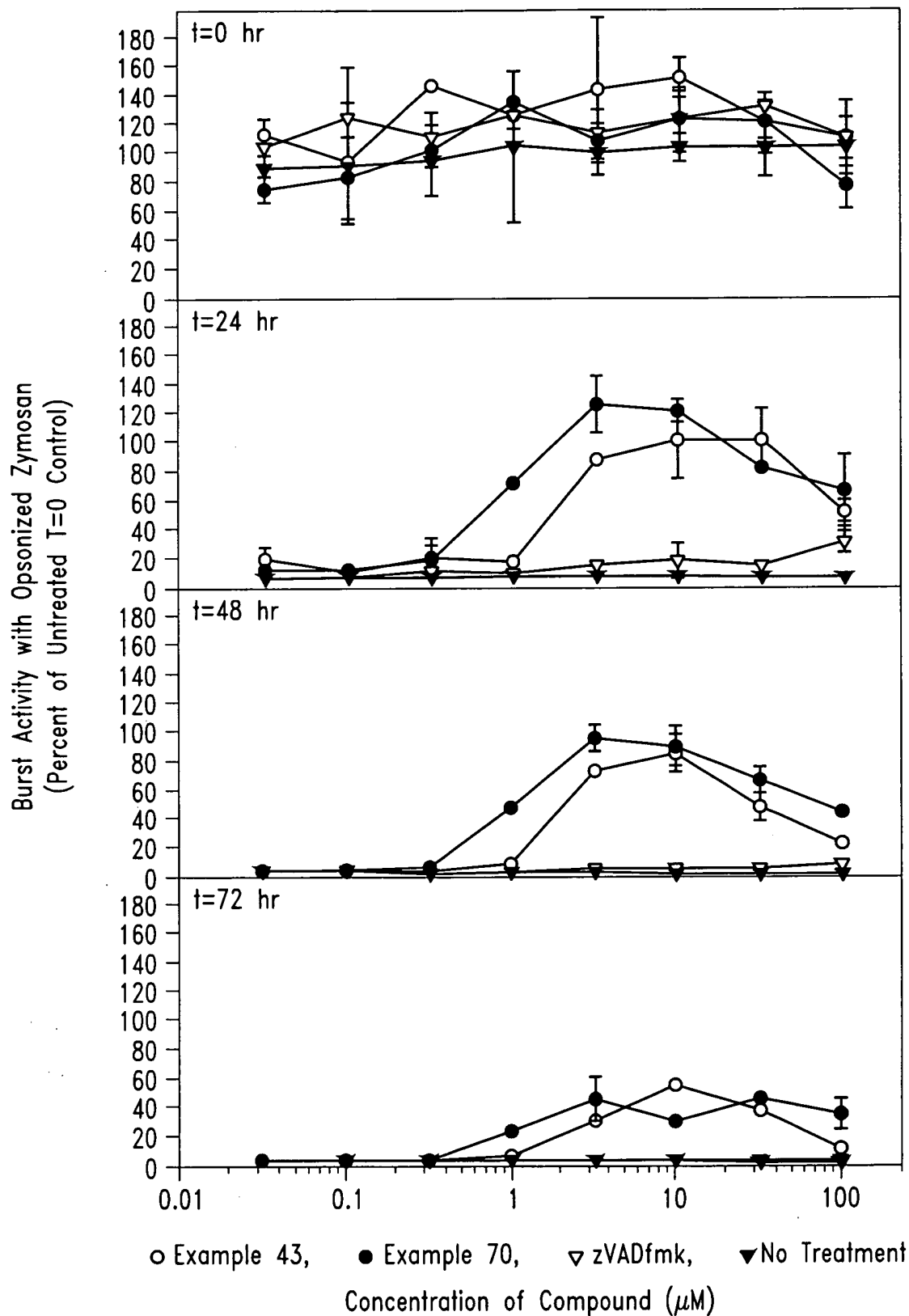


Fig. 7